## REMARKS/ARGUMENTS

Reconsideration of this patent application is respectfully requested in view of the foregoing amendments, and the following remarks. Claims 1-8 are in the application. Claims 3, 6 and 7 have been amended. The specification and sequence listing have been amended. No new matter has been added.

The Examiner requested that a typed version of Form 1449 from the Information Disclosure Statement be submitted.

Applicants submit herewith a typed version of the form, containing a listing of the references already submitted.

The Examiner objected to the specification for lacking headings and a brief description of FIG. 1. Applicants have amended the specification to add headings and to include a description of FIG. 1.

The Examiner objected to the sequence listing, stating that the CRF copy submitted is flawed. Applicants submit herewith a corrected CRF copy of the sequence listing, and the required statement.

The Examiner objected to claim 3 for referring to a Table.

Applicants have amended claim 3 in accordance with the Examiner's suggestions, to include a listing of the specific mutations from the table.

The Examiner rejected claims 6-8 under 35 U.S.C. §112, stating that the plasmid of claims 6-8 must be obtainable or available. Applicants submit that a deposit of the plasmid of claims 6-8 was not made because the plasmid is easily reproducible by one of skill in the art. The plasmid of claim 6 can be easily reproduced because the only difference to a known strain is the presence of a metA allele of claim 4. allele is clearly defined in the specification and in claim 4. One of skill in the art can easily make this metA allele, as well as the plasmid containing this metA allele (claim 5), as well as the isolated microbial cell of claims 6 and 7 using standard procedures that have been well known for many years. examples describe the procedures for the production of several different plasmids, as well as the production of microorganisms using these plasmids (Examples 2 and 3). Therefore, a deposit according to the Budapest treaty is unnecessary for the invention.

The Examiner also rejected claims 6-8 because it was unclear which strain was required. The Examiner suggested amending claims 6-8 to recite "An isolated microbial host cell." Applicants have amended the claims accordingly.

The Examiner has rejected claims 1-8 under 35 U.S.C. §112, first paragraph, as not being sufficiently described in the specification. The Examiner also rejected claims 1-8 under 35 U.S.C. §112, first paragraph, for not being enabled.

Applicants respectfully traverse.

The enzyme of claim 1 is defined by the mutation of Asp and Tyr at conserved sites of the protein plus a functional definition (i.e. the reduced sensitivity towards L-methionine or SAM). Some mutants of the wild-type enzyme and the methods to determine the feedback resistance are provided in the description wherein the conserved sites are replaced by various amine acids. Therefore, the scope of claim 1 is appropriate because a person skilled in the art can recognize that the mutants claimed in claim 1 achieve the purpose of the present application.

Therefore, the sequences/proteins according to the present invention can be made by a person skilled in the art using the information of the present specification, and the examples of the present application prove the interrelation of the sequence and the desired features. Enclosed as Appendix A is an alignment of the SAM sequences of 14 different species. This alignment shows that in all these enzymes, Asp in position 101 and Tyr in Position 294 (both marked green) are conserved. In the light of this state of the art it is unnecessary to limit claim 1 to SEQ. ID. No.: 2, as claim 1 as written is in compliance with 35 USC §112.

The Examiner rejected claims 4 and 5 under 35 U.S.C. §101 for non-statutory subject matter, and suggested that claims 4 and 5 be amended to claim "an isolated nucleic acid encoding homoserine transsuccinylase". Applicants have amended claims 4 and 5 accordingly. Applicants submit that claims 4 and 5 are now in compliance with 35 U.S.C. §101.

The Examiner has rejected claims 1-8 under the judicially-created doctrine of double patenting over claims 1-8 of copending Application Serial No. 10/530,844. Applicants submit herewith a

Terminal Disclaimer, disclaiming that portion of any patent issuing on the present application that would extend beyond the term of the patent issuing from application serial no. 10/530,844.

Accordingly, Applicants submit that the application is in condition for allowance. Early allowance of the claims is respectfully requested.

Respectfully submitted,

Susanme LEONHARTSBERGER ET AL.

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ECR: cmm

Enclosure:

Appendix A

Corrected Sequence Listing, Disk and Statement

Terminal Disclaimer and fee Typed version of PTO Form 1449

I hereby certify that this correspondence is being deposited with the U.S. Postal Service as first class mail in an envelope addressed to: Commissioner of Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on May 24, 2007.

85/31/2007 GFREY1

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## Appendix A

MRIKIPDNLPAKETLTKENIFVMAESRAYSODIRPLKIVILNIMEIKOTT MPINIPTHLPAKOVIESEHIFVMDESRAFHÖDIRPOKITILNIMPKKIOT MPIKIPDDLPATSVIEAEGVMVMREADAVRODIRPLKIGILNIMPKKVTT MPIKIPDNLPAAKTINEENIFFMDEDRAYHODIRPLNIVIVNIMPTKIVM MetA\_B.halodurans (1)MetA\_B.subtilis (1)MetA\_Bruc.meritensis (1)MetA\_C.acetobutylicum (1) MetA\_Camp.jejuni MetA\_E.coli MetA\_L.lactis MPLIIPENIPAYELLK-EHAFIMGLRRAKHODIRPOEILIVNLMPKKIET METRVPDELPÄVNFLREENVFVMTTSRASGOETRPLKVLILNIMPKKIET MPVKVIEGLPATDDLRADNIFVMNDERAKNONTRPLNILVVNIMPRKLIT (1) (1) MPIKIPDTLPAFETLVHEGVRIMTETEAIRQDIRPLQIGLLNIMPNKIKT (1) MetA\_R.meliloti MEIRVLDELPÄVNFIREENVFVMTTSRÅSGÖEIRPLKVLILNIMPKKIET MEIRVLDELPÄVNFIREENVFVMTTSRÅSGÖEIRPLKVLILNIMPKKIET MEIRIDKKLPÄVEIIRTENIFVMDDORÅAHQDIRPLKILILNIMPKKIET MEINVPSGLPÄVKVIAKEGIFVMTEKRÄIHQDIRPLEILILNIMPKKIKT (1) MetA\_S.typhi MetA\_S.typhimurium (1)MetA\_Str.pneumoniae (1)MetA\_T.maritima (1) metrtpdolpasdylrnenifymsesrastoetrelkylilnimekkiet metrypdelpaysfirnenyfymassraktoetrelkylilnimekkiet MetA\_V.cholerae (1) MetA\_Y.pestis MPIRIPD LPAV VLR ENIFVM ESRA QDIRPLKILILNLMPKKI T Consensus (1)ETQLLRLLGNTPLQVEVSFMYTDTHISKNTSYDHLQTFYQTIDEVKQKKF MetA\_B.halodurans (51)ETOLLRLIGNTPLQVEVSFMYTDTHISKNTSYDHLQTFYQTIDEVKQKKF ETOLLRLIGNSPLQVHTTFLIPSTHTPKNTAREHLDEFYTTFSNTRHKRF ETOLARLIGATPLQVELTLVRMTNHVARHTPADHMLSFYCPWEEVNDQRF ETOLLRLIGNSPLQVNPTFIHTQTHKSQNTSKEHLIKFYETFEELKNNKF ENOLLSLLANSPLQVNITLLATTSYVGKNTPFTHLEKFYKGLEEVKKHKF ENOFLRLISNSPLQVDIQLLRIDSRESRNTPAEHLNNFYCNFEDIQDQNF ERQTLRLISNTPLQTNVEFLYMTSHDFKNTKQGHLDSFYKSFSEIKSQYY EIOMARLIGATPLQVEITLVRVNGHRPKNTPEEHLNFYCTFEEVEARKF ENOFLRLISNSPLQVDIOLLRIDARESRNTPAEHLNNFYCNFDDTCDONF (51) MetA\_B.subtilis \_Bruc.meritensis (51)MetA\_C.acetobutylicum (51)MetA\_Camp.jejuni MetA\_E.coli (50) (51)MetA\_L.lactis (51)(51)MetA\_R.meliloti ENOFLRLLSNSPLOVDIQLLRIDARESRNTPAEHLNNFYCNFDDICDQNF MetA\_S.typhi (51)MetA\_S.typhimurium (51) ENOFLELISMSPLOVDIQLLRIDARESENTPAEHLNNEYCNFDDICDQNF (51) ETOLLRHLANTPLOLDIDFLYMESHRSKTTRSEHMETFYKTFPEVKDEYF MetA\_Str.pneumoniae MetA\_T.maritima MetA\_V.cholerae EIOLLRLIGHTPLOVNVTLLYTETHKPKHTPIEHILKFYTTFSAVKDRKF (51)ETOFLELLSNSPLOVDIELLRIDDRPSKNTPEEHLNTFYRQFELVKNRNF (51) ENOFLRLLSNSPLOVDIQLLRYDSRESKNTPTEHLNNFYCDFEDIQDQNF MetA\_Y.pestis (51) E QILRLISNSPLQVDITLLRIDSH SKNTP EHL FY TFEEIKDQKF Consensus (51)GMITTGAPTETLPYDEVDYWNELKQIMEWSKTNVTSTLHICWGAQAGLF MetA\_B.halodurans (101)D.MIITGAPIEHLAFEEVSYWEELKEIMEWSKTNYTSTLHICWGAQAGLY MetA\_B.subtilis (101)FVITGAPVERLPFEEYTYWDEMRRYFDWTQSHVHRTLNICWAAQAAVY (101) MetA\_Bruc.meritensis MIVTGAPVETLSFENVDYWEELCRIFDWSVTNVTSTIHICWGAQAGLY (101)MetA\_C.acetobutylicum MetA\_Camp.jejuni MetA\_E.coli AIVTGAPVEOMDFEKVAYWEELLEIFDFLKONVTSSMYICWGAMAALK (100)LIVTGAPIGLVEFNOVAYWPOIKOVLEWSKOHVTSTLFVCWAVQAALN
LIVTGAPVEQLNFEEVDYWSELLKIIDWSKSHVYSSLHICWGAQAALY (101)(101) MetA\_L.lactis FITTGAPIETLEYEEYTYWKELQRIFDWTTTNVHSTLNVCWGGMAAVY (101) MetA\_R.meliloti (101) FITGAPIETLEYEEVTYWKELQRIFDWTTTNVHSTLNVCWGMAAVY
(101)
(101) LIVTGAPIGLVEFNDVAYWPQIRQVLEWAKDHVTTSTLFVCWAVQAALN
(101) LIVTGAPIGLVEFNDVAYWPQIRQVLEWAKDHVTSTLFVCWAVQAALN
(101) MITGAPVEHLPFEEVDYWEEFRQMLEWSKTHVYSTLHICWGAQAGLY
(101) LITGAPUGLVQFEDVAYWQHLQNIMAWAKAHVTSTLYICWAAQAGLK
(101) LITGAPUGLVQFEDVAYWQHLQNIMAWAKAHVTSTLYICWAAQAGLK
(101) LITGAPUGLVDFCDVAYWPQIERITAWAKEHVTSTLFVCWAVQALN
(101) LITGAPVE L FEEVAYW ELR ILEWSKTHVTSTLHICWAAQAALY MetA\_S.typhi MetA\_S.typhimurium MetA\_Str.pneumoniae MetA\_T.maritima MetA\_V.cholerae MetA Y.pestis Consensus 151 YHYGVEKVPLPEKOFGVYPHKINVPNVKLLRGFDDEFYVPHSRHTDINKA MetA\_B.halodurans YHYGVEKIOMPKKIFGVFEHTVLSKHERLVRGFDELYYVFHSRHTDINME HFHGWKYDLPAKASGVFRQRSLVLASPYLRGFSDDFALEVSRWTEVRKS MetA\_B.subtilis (151)(151)MetA\_Bruc.meritensis HHYGIPKYELHEKLFGVFKHNLTERNIKLTRGFDDEFYAPHSRHTYVKRE YFYGVDKISLDKKIFGVYKHDKVSPDLLLTN-LDEKVLMPHSRHSSMDEE ILYGIPKOTRTEKLSGVYEHHILHPHALLTRGFDDSFLAPHSRYADFPAA MetA\_C.acetobutylicum (151)MetA\_Camp.jejuni (150)MetA\_E.coli (151)ARYGYTKENLPOKLCGIYKSSVEQPKNPLFRGFDDFFNYPQSRYTQSNPS HFHGYPKYPLKEKAFGVYRHONLQPSSVYLNGFSDDFAVEVSRWTEVRRA MetA\_L.lactis (151)MetA\_R.meliloti (151)ILYGJPKOTRTDKLSGVYEHHILHPHALLTRGFDDSFLAPHSRYADFPAA (151)MetA\_S.typhi ILYGIPKOTRTDKLSGVYEHHILHPHALLTRGFDDSFLAPHSRYADFPAA MetA\_S.typhimurium (151)MetA\_Str.pneumoniae
MetA\_T.maritima
MetA\_V.cholerae
MetA\_Y.pestis LRYGVEKYQMDSKLSGIYPQDTLKEGHLLFRGFDDSYVSPHSRHTEISKE (151)YFYGIPKYELPOKLSGVYKHR-VAKDSVLFRGHDDFFWAPHSRYTEVKKE (151)(151)LLYNLPKRTREEKLSGVYYHDIHKPFHPLLRGFDDRFLAPHSRYADFDAE ILYGIPKMTREVKLSGIYQHQTLEPLALLTRGFDETFLAPHSRYADFPVE (151)(151) Y YGIPK L EKLSGVY H IL P ALL RGFDD FLAPHSRYTDV Consensus QIEAHPDLEITSESEQ-AGYYTVASKOGKQIFVTGHSEYDACTTQQEYER (201)MetA\_B.halodurans OLOAVPELNILTASKEAGGLLLIVSKDEKOVFLTGHPEYDTNTLLOEYER DIPADSGLKYLVDSTE-TGLCLLDDPRHRSLHMPNHVEYDTTSHADEYFR MetA\_B.subtilis (201)(201) MetA\_Bruc.meritensis MetA\_C.acetobutylicum
MetA\_Camp.jejuni
MetA\_E.coli
MetA\_L.lactis DIKKNPSLKILAESDE-AGAYIVASENGKNIFVMGHAEYDGDTLNLEYIR (201)QILALQKQGKLKILLRNKKIGSALLRDEKNIFILGHLEYFKETLHQEYVR (199)LIRDYTDLEILÄETEE-GDAYLFASKDKRIAFVTGHPEYDAQTLAQEFFR (201)EIKKVPDLEVLSSSKE-TGFSILAKKNLREIYLFGHLEYDRETLAWEYER (201)(201)DIDRVPDLEILMESKE-VGVCLVHEKKGNRLYMFNHVEYDSTSLSEEYFR MetA\_R.meliloti LIRDYTDLEILAETEE-GDAYLFASKDKRIAFVTGHPEYDAHTLAGEYFR (201) MetA\_S.typhi (201) LIRDYTDLEILAETEE-GDAYLFASKDKRIAFVTGHPEYDAHTLAGEYFR MetA\_S.typhimurium

MetA\_Str.pneumoniae (201) EVLNKTNLEILSEGPQ-VGVSILASRDLREIYSFGHLEYDRDTLAKEYFR MetA\_T.maritima (200)DIDKVPELEILAESDE-AGVYVVANKSERQIFVTGHPEYDRYTLRDEYYR MetA\_V.cholerae (201) FLAEHTDLDILATSDV-AGVYLAATKDKRNVFVTGHPEYDAYTLHGEYVR MetA\_Y.pestis (201)VLQQYTDLDILVSSEE-AGAYLFASKDKRVAFVTGHPEYDVDTLAGEYQR Consensus (201)TDLEILAESEE AGVYLVASKD R IFVTGHPEYD TLA EYFR (250)DRAR-GLNIQVPENYFPNDDATRKPLLRWRAHSYLLFSNWLNYYVI ETP DLERNLSTVEAPKHYFAKGSNE--PVNRWKAHATLLFMNWLNYYVY ETP MetA\_B.halodurans MetA\_B.subtilis (251)DIQV-QPEAKVPVNYFPGDDAKRPPENRWRSHAHLLFGNWIN-EM MetA\_Bruc.meritensis (250)MetA\_C.acetobutylicum (250)DKNQ-GMNIKIPKNYFKDNDPEKGPMVTWRGHANLLFSNWLNYYV ETP MetA\_Camp.jejuni (249)-D--N-FIQKAKNYYDKKGN---IKYNWRSNANTIFANWLNYD MetA\_E.coli (250)DVEA-GLDPDVPYNYFPHNDPQNTPRASWRSHGNLLFTNWLNYY MetA\_L.lactis DKEE-GLKPNLPQNYFPENDDKNKPKSTWASAASLFFSNWLNYAV (250)DVDA-GVPIKLPHDYFPHNDSALPPQNRWRSHAHLFFGNWIN-ET TTP MetA R meliloti (250)MetA\_S.typhi (250)DVEA-GLNPEIPYNYFPKNDPQSIPRTTWRSHGNLLFTNWLNYYW DITP MetA\_S.typhimurium (250)DVEA-GLNPEVPYNYFPKNDPQNIPRATWRSHGNLLFTNWLNYY CLTP MetA\_Str.pneumoniae (250)DRDA-GFDPHIPENYFKDDDVNQVPCLCWSSSAALFFSNWVDHAW: ETP MetA\_T.maritima (249)DIGR-NLKVPIPANYFPNDDPTKTPILTWWSHAHLFFSNWLNYC MetA\_V.cholerae (250)DLGE-GLNPAIPVNYYPNDNPDNKPCASWRSHGHLLFANWLNYCW DQTP DLAA-GLNPQVPLNYFPSDDASLRPKASWRSHGHLLFANWLNYW 10 ITP DVE GL P IP NYFP D P TWRSHA LLFSNWLNYY 1 TP MetA\_Y.pestis (250)Consensus (251)301 (299)YDLSR-----MetA\_B.halodurans (299)YEWD-----MetA\_B.subtilis (298) YDIERIGKV-----MetA\_Bruc.meritensis (299) FEL-----MetA\_C.acetobutylicum FVL-----MetA\_Camp.jejuni (291)(299)YDLRHMNPTLD-----MetA\_E.coli MetA\_L.lactis YLGERLSQHLNEENYDFNQKEQK (299)YELAKIGTGER-----MetA R.meliloti (298)YDLRHMNPTLD-----MetA\_S.typhi (299)MetA\_S.typhimurium (299) YDLRHMNPTLD-----(299) FDWRKIEDDASAYGYL-----MetA\_Str.pneumoniae YRLEDIH-----MetA\_T.maritima (298)(299) YDLEKFSEANFTKDE-----MetA\_V.cholerae MetA\_Y.pestis (299) FDLRHMNPTLD-----

Consensus

(301) YDL